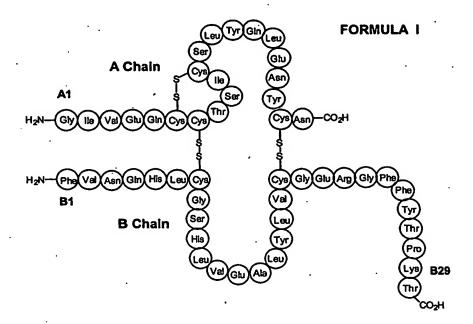
## WHAT IS CLAIMED IS:

- 1. An insulin derivative comprising an insulin molecule and a reactive group for covalently bonding a blood component.
- 2. The insulin derivative of claim 1, wherein the insulin molecule is of formula I:

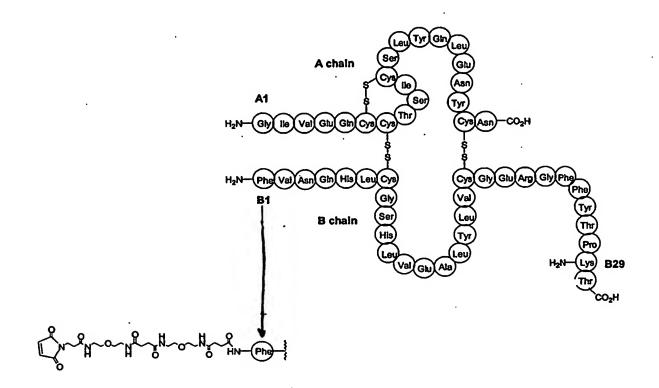


and the reactive group is coupled to an amino acid of the insulin molecule at a position selected from the positions Gly A1, Phe B1 and Lys B29.

- 3. The insulin derivative of claim 1 or 2, wherein the reactive group is selected from the group consisting of Michael acceptor, a succinimidyl-containing group and a maleimido-containing group.
- 4. The insulin derivative of claim 3, wherein the Michael acceptor is an  $\alpha,\beta$ , unsaturated carbonyl moiety.
- 5. The insulin derivative of claim 1 or 2, wherein the reactive group is a maleimido-containing group.

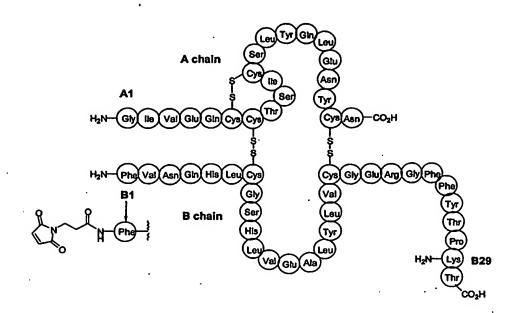
- 6. The insulin derivative of claim 1 or 2, wherein the reactive group is 3-Maleimidopropionic acid (MPA).
- 7. The insulin derivative of any one of claims 1 to 6, wherein the reactive group is coupled to an amino acid of the insulin molecule via a linker.
- 8. The insulin derivative of claim 7, wherein said linker is selected from the group consisting of (2-amino) ethoxy acetic acid (AEA), ethylenediamine (EDA), amino ethoxy ethoxy succinimic acid (AEES), AEES-AEES, 2-[2-(2-amino)ethoxy)] ethoxy acetic acid (AEEA), AEEA-AEEA, -NH<sub>2</sub>-(CH<sub>2</sub>)<sub>n</sub>-COOH where n is an integer between 1 and 20 and alkyl chain (C1-C10) motif and combination thereof.
- 9. The insulin derivative of claim 8, wherein said alkyl chain (C1-C10) motif is one or more alkyl chains (C1-C10) saturated or unsaturated in which could be incorporated oxygen nitrogen or sulfur atoms.
- 10. The insulin derivative of claim 9, wherein said alkyl chain is selected from the group consisting of glycine, 3-aminopropionic acid (APA), 8-aminooctanoic acid (AOA) and 4-aminobenzoic acid (APhA).
- 11. The insulin derivative of claim 8, wherein said combination is selected from the group consisting of AEEA-EDA, AEEA-AEEA and AEA-AEEA.
- 12. The insulin derivative of claim 6, wherein said linker is  $-NH_2-(CH_2)_7-COOH$ .

13. The insulin derivative of claim 1 having the formula:



## 14. The insulin derivative of claim 1, having the formula:

15. The insulin derivative of claim 1, having the formula:



- 16. The insulin derivative of claim 1, wherein said blood component is a blood protein.
- 17. The Insulin derivative of claim 16, wherein said blood protein is serum albumin.
- 18. An Insulin conjugate comprising an insulin derivative according to any one of claims 1 to 17 and a blood component, wherein the reactive group and the blood component are conjugated through a covalent bond formed between said reactive group and said blood component.
- 19. The insulin conjugate of claim 18, wherein the blood component is a blood protein.
- 20. The insulin conjugate of claim 19, wherein the blood protein is serum albumin.

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- 21. The insulin conjugate of claim 18, wherein said conjugate was formed ex vivo.
- 22. A pharmaceutical composition comprising the insulin derivative of any one of claims 1 to 17 in association with a pharmaceutically acceptable carrier.
- 23. A pharmaceutical composition comprising the insulin conjugate of any one of claims 18 to 21 in association with a pharmaceutically acceptable carrier.
- 24. A method for treating a glycaemic-related disease or disorder in a subject suffering from said glycaemic-related disease or disorder, comprising administering the insulin derivative of any one of claims 1 to 17 to said subject.
- 25. The method according to claim 24, wherein said glycaemic-related disease is selected from the group consisting of diabetes of type I,

diabetes of type II, gestational diabetes, cystic fibrosis, polycystic ovary syndrome and pancreatitis.

- 26. The method according to claim 24, wherein the glycaemic-related disease is selected from the group consisting of diabetes of type I and diabetes of type II.
- 27. A method for treating a glycaemic-related disease or disorder, comprising the administration of the insulin conjugate of any one of claims 18 to 21.
- 28. The method according to claim 27, wherein said glycaemic-related disease is selected from the group consisting of diabetes of type I, diabetes of type II, gestational diabetes, cystic fibrosis, polycystic ovary syndrome and pancreatitis.
- 29. The method according to claim 27, wherein the glycaemic-related disease is selected from the group consisting of diabetes of type I and diabetes of type II.
- 30. A method for treating a glycaemic-related disease or disorder, comprising the administration of the pharmaceutical composition of any one of claims 22 and 23.
- 31. The method according to claim 30, wherein said glycaemic-related disease is selected from the group consisting of diabetes of type I, diabetes of type II, gestational diabetes, cystic fibrosis, polycystic ovary syndrome and pancreatitis.
- 32. The method according to claim 30, wherein the glycaemic-related disease is selected from the group consisting of diabetes of type I and diabetes of type II.
- 33. Use of the derivative of any one of claims 1 to 17, for the preparation of a medicament for the treatment of a glycaemic-related disease or disorder.

- 34. The use as claimed in claim 33, wherein said glycaemic-related disease is selected from the group consisting of diabetes of type I, diabetes of type II, gestational diabetes, cystic fibrosis, polycystic ovary syndrome and pancreatitis.
- 35. The use as claimed in claim 33, wherein the glycaemic-related disease is selected from the group consisting of diabetes of type I and diabetes of type II.
- 36. Use of the conjugate of any one of claims 18 to 19, for the preparation of a medicament for the treatment of a glycaemic-related disease or disorder.
- 37. The use as claimed in claim 36, wherein said glycaemic-related disease is selected from the group consisting of diabetes of type I, diabetes of type II, gestational diabetes, cystic fibrosis, polycystic ovary syndrome and pancreatitis.
- 38. The use as claimed in claim 37, wherein the glycaemic-related disease is selected from the group consisting of diabetes of type I and diabetes of type II.